1)9 620,820 PTO/SB/06 (08-03)

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U.S. Patent and Trademark Office; U.S. DEPARTMENT OF COMMERCE

Under the Paperwork Reduction Act of 1995, no persons are required to respond to a collection of information unless it displays a valid OMB control number. PATENT APPLICATION FEE DETERMINATION RECORD **Application or Docket Number** Substitute for Form PTO-875 CLAIMS AS FILED - PART I OTHER THAN OR SMALL ENTITY SMALL ENTITY (Column 1) (Column 2) FOR NUMBER FILED NUMBER EXTRA RATE FEE RATE FEE BASIC FEE (37 CFR 1.16(a)) OR TOTAL CLAIMS (37 CFR 1.16(c)) minus 20 = X S OR INDEPENDENT CLAIMS = (37 CFR 1.16(b)) minus 3 = X S OR X \$ MULTIPLE DEPENDENT CLAIM PRESENT (37 CFR 1.16(d)) OR * If the difference in column 1 is less than zero, enter "0" in column 2. TOTAL OR TOTAL CLAIMS AS AMENDED - PART II OTHER THAN OR (Column 2) (Column 3) SMALL ENTITY (Cotumn 1) SMALL ENTITY CLAIMS HIGHEST PRESENT REMAINING NUMBER RATE RATE ADDI-**EXTRA** ENDMENT **AFTER PREVIOUSLY** TIONA TIONAL AMENDMENT PAID FOR FF FEE Total (37 CFR 1.16(e)) Minus OR X S Independent (37 CFR 1.16(b)) Minus 2 OR FIRST PRESENTATION OF MULTIPLE DEPENDENT CLAIM (37 CFR 1.16(d)) OR TOTAL TOTAL OR ADD'L FEE ADD'L FEE (Calumn 2) (Column 3) (Column 1) CLAIMS HIGHEST PRESENT NUMBER RATE RATE REMAINING ADDI. ADDI-AMENDMENT PREVIOUSLY TIONAL TIONAL AFTER FEE MENDMENT PAID FOR FEE Total Minus (37 CFR 1.16(c)) OR Independent (37 CFR 1.16(b)) Minus Юd x \$ 100 = OR X \$ FIRST PRESENTATION OF MULTIPLE DEPENDENT CLAIM (37 CFR 1.18(d)) OR TOTAL TOTAL 'n∂ ADD'L FEE OR ADD'L FEE (Column 1) (Column 2) (Column 3) HIGHEST CLAIMS O PRESENT REMAINING NUMBER RATE ADDI-RATE ADDI: PREVIOUSLY **EXTRA** TIONAL TIONAL ENDMENT AFTER AMENDMENT PAID FOR FEE FEE Total Minus (37 CFR L18(c)) X S OR Minus independent (37 CFR 1.16(b)) X S = X S OR FIRST PRESENTATION OF MULTIPLE DEPENDENT CLAIM (37 CFR 1.16(d)) OR TOTAL TOTAL ADD'L FEE OR ADD'L FEE If the entry in column 1 is less than the entry in column 2, write "0" in column 3. ** If the "Highest Number Previously Paid For" IN THIS SPACE is less than 20, enter "20". "If the "Highest Number Previously Paid For" IN THIS SPACE is less than 3, enter "2".

"If the "Highest Number Previously Paid For" IN THIS SPACE is less than 3, enter "2".

The "Highest Number Previously Paid For" (Total or Independent) is the highest number found in the appropriate box in column 1.

This collection of information is required by 37 CFR 1.16. The information is required to obtain or retain a benefit by the public which is to file (and by the USPTO to process) an application. Confidentiality is governed by 35 U.S.C. 122 and 37 CFR 1.14. This collection is estimated to take 12 minutes to complete, including gathering, preparing, and submitting the completed application form to the USPTO. Time will vary depending upon the individual case. Any comments on the amount of time you require to complete this form and/or suggestions for reducing this burden, should be sent to the Chief Information Officer, U.S. Petent and Trademark Office, U.S. Department of Commerce, P.O. Box 1450, Alexandria, VA 22313-1450. DO NOT SEND FEES OR COMPLETED FORMS TO THIS ADDRESS. SEND TO: Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450.

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eby certify that this correspondence is being deposited with the United States Postal Service on the date set forth below as First Class Mail in an envelope addressed to: Mail Stop Amendment, Commissioner for Patents, P.O. Box 1450, Alexandria VA

Date of Signature and Deposit: May 17, 2006

PATENT

Vinarov

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicants:

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Date: May 17, 2006

Donald L. Gillian-Daniel

Paul W. Bates

Serial No.: 09/620,820

Group Art Unit: 1636

Filed: July 21, 2000

Examiner: Celine X. Qian

Title: INHIBITION OF LIPOPROTEIN SECRETION

File No.: 960296.97290

RESPONSE

MAIL STOP AMENDMENT **Commissioner For Patents** P. O. Box 1450 Alexandria, VA 22313-1450

Dear Sir:

In response to the Office Action dated January 23, 2006, the applicants respond herewith as follows:

Amendments to the Claims are reflected in the listing of claims which begins on page 2 of this paper.

Remarks begins on page 6 of this paper.

Application No.: 09/620,820 Amendment dated May 17, 2006 Reply to Office Action of January 23, 2006

domains for the ER were known before, whether or not this level of localization would be sufficient to locate the LDL receptor, where it could interact with apoB, was unknown. It was not clear at all if the ER was the right place in the secretion pathway for trapping apoB. Despite, any lack of suggestion for the literature, applicants identified an inventive route of delivery and demonstrated that it was effective in lowering serum cholesterol in mammals. This was a highly specific result.

In summary, applicants submit that the combination of the uncertainty arising in general with techniques of genetic engineering, and the uncertainty inherent in dealing with a complex biological system such as cholesterol secretion, makes it unpredictable that the method described by the applicants here would actually work. Thus, there was no motivation or reasonable expectation that combining the cited documents would be successful in advance of applicants' claimed invention.

Accordingly, applicants respectfully request that in view of the supplemental Declaration and comments, the rejection be respectfully reconsidered and withdrawn, and that a timely Notice of Allowance be issued in this case.

A petition for a one-month extension of time is enclosed. No other fees are believed due in regard to this submission. If any other fee is due or any other extension of time is required in this or any subsequent response, please consider this to be a petition for the appropriate extension and a request to charge the petition fee to the Deposit Account No. 17-0055.

Respectfully submitted,

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